

#### (1) GENERAL INFORMATION:

- (i) APPLICANT: TSYRLOVA, IFENA WOLPE, STEPHEN D.
- (ii) TITLE OF INVENTION: INHIBITOR OF STEM CELL PROLIFERATION AND USES THEREOF
- (111) NUMBER OF SEQUENCES: 37
- (iv) CORRESPONDENCE ADDRESS:
  - (A) ADDRESSEE: NIMON & VANDERHYE P.C.
  - (B) STREET: 1100 FORTH GLEER ROAD
  - (C) CITY: ARLINGTON
  - (D) STATE: WIRGINIA
  - (E) COUNTRY: U.S.A.
  - (F) DIF: 20001-4704
  - (v) COMPUTER READABLE FORM:
    - (A) MEDIUM TYPE: Floppy disk
    - (B) COMPUTER: IBM PC compatible
    - (C) OPERATING SYSTEM: PC-DOS/MS-DOS
    - (D) SOFTWARE: PatentIr. Release #1.0, Version #1.30
- (v1) CURRENT APPLACATION DATA:
  - (A) APPLICATION NUMBER: US 08:832,443
  - (B) FILING DATE: Un-APR-1997
  - (C) CLASSIFICATION:
- (viii) ATTORNEY AGENT INFORMATION:
  - (A) NAME: BYRNE, THOMAS E.
  - (B) REGISTRATION NUMBER: 32,205
  - (C) REFERENCE/DOCKET NUMBER: 1331-222
  - (ix) TELECOMMUNICATION INFORMATION:
    - (A) TELEPHONE: (70.) 816-4000
    - (3) TELEFAM: (103) 916-4100
  - THA FMATT OF FOR BOY ID NOTE:
    - FOR SEQUENCE CHARACTERISTICS:
      - (A) LENGTH: 13 amino acids
      - (B) FYPE: amino acid
      - (C) STRANDEDNESS:
      - pr Topology linear
    - on the first paper. See 1 de
    - an language Ang Leu Jer Hit Ly Ala Asi Asi Asi

(B) TYPE: amino acid (C) STRANDEDNESS: (D) TOPOLOGY: linear (ii) MOLECULE TYPE: peptide (x1) SEQUENCE DESCRIPTION: SEQ ID NO:2: Cys Phe Pro His Phe Asp Leu Ser His Gly Ser Ala Gln Val Cys 10 (2) INFORMATION FOR SEQ ID NO:3: (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 19 amino acids (E) TYPE: amino acid (C) STRANDEDNESS: (D) TOPOLOGY: linear (11) MOLECULE TYPE: peptide (xi) SEQUENCE DESCRIPTION: SEQ ID NO:3: Asp Ala Leu Thr Asn Ala Val Ala His Val Asp Asp Met Pro Asn Ala 10 Leu Ser Ala 10: INFORMATION FOR SEQ ID NO:4: . SEQUENCE CHARACTERISTICS: At LEN Net: le arin catilis AB TYPE: amino atrib (D) TOPOLOGY: linear - (i) MOLECULE TYPE: peptide

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(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 15 amino acids

- (C) STRANDEDNESS:
- (D) TOPOLOGY, linear
- Will MOLECULE TYPE, peptide
- (xi) SEQUENCE DESCRIPTION: SEQ ID NO:5:

Leu Val Val Tyr Pro Trp Thr Gln Arg 1 5

- (2) INFORMATION FOR SEQ ID NO:6:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 8 amine acids
    - (B) TYPE: amino acid
    - (C) STRANDEDNESS:
    - (D) TOPOLOGY: linear
  - (ii) MOLECULE TYPE: peptide
  - (xi) SEQUENCE DESCRIPTION: SEQ ID NO:6:

Leu Val Val Tyr Pro Trp Thr Gln 1

- (2) INFORMATION FOR SEQ ID NO:7:
  - (i) SEQUENCE CHAFACTERISTICS:
    - (A) LENGTH: " amino acids
    - (B) TYPE: amino acid
    - (C) STRANDEDNESS:
    - (D) TOPOLOGY: linear
  - The MOLECULE TYPE peptide
  - (xi) SEQUENCE DESCRIPTION: SEQ ID No:2:

led Val Val Tyr Pro Trp Thr

THE EMAID NOTES SE, IN THE

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:8: Leu Val Val Tyr Pro Trp (2) INFORMATION FOR SEQ ID NO:9: (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 5 amino acids (B) TYPE: amino acid (C) STRANDEDNESS: (D) TOPOLOGY: linear (ii) MODECULE TYPE, peptide (xi) SEQUENCE DESCRIPTION: SEQ ID NO:9: Leu Val Val Tyr Pro (2) INFORMATION FOR SEQ ID NO:10: (i) SEQUENCE CHAFACTERISTICS: (A) LENGTH: " amino acids (B) TYPE: amino acid (0) STRANDEDNESS: (D) TOPOLOGY: linear (ii) MOLECULE TYPE: peptide (xi) SEQUENCE DESCRIPTION: SEQ ID NO:10: Chai tal the PESS Tep The Win in amainn bu st. in naide (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 7 amino acids (B) TYPE: amino acid (C) STRANDEDNESS: (D) TOPOLOGY: linear ii Mi Paul Paul Paul Peptide lyr e. By Incom Arche

(1) SEQUENCE CHARACTERISTICS: (A) LENGTH: 6 amino acids (B) TYPE: amino acid (C) STRANDEDNESS: (D) TOPOLOGY: linear (ii) MODECULE TYPE: peptide (xi) SEQUENCE DESCRIPTION: SEQ 1D NO:12: Tyr Pro Trp Thr Gln Arg (2) INFORMATION FOR SEQ ID NO:13: (i) SEQUENCE CHAFACTERISTICS: (A) LENGTH: 5 amino acids (B) TYPE: amino acid (C) STRANDEDNESS: (D) TOPOLOGY: linear (ii) MOLECULE TYPE: peptide (xi) SEQUENCE DESCRIPTION: SEQ ID NO:13: Tyr Pro Trp Ihr Gln (2) INFORMATION FOR SEQ ID NO:14: (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 6 amino acids (B) TYPE: amino acid (C) STRANDEDNESS: an: ToPOLOGY: linear I MMETHE TYPE pertibe \*X: SEQUENCE DESCRIPTION: SEQ ID NO: 14: The Mor Ten Mor Pho Arm

A TABLET CONFIDENCE OF A TABLE OF

# (ii) MOLECULE TYPE: DNA (genomic)

	(xi) SI	EQUENCE DESC	CRIPTION: SE	EQ ID NO:15:			
GT GC	TGICIC	CTGCCGACAA	GACCAACG1C	AAGGCCGCCT	GGGGTAAGGT	CGGCGCGCAC	+5 C
GCTC	GCGAGT	ATGGTGCGGA	GGCCCTGGAG	AGGATGTTCC	TGTCCTTCCC	CACCACCAAG	1: C
ACC.	ACTICC	CGCACTTCGA	CCTGAGCCAC	GGCTCTGCCC	AGGTTAMGGG	CCACGGCAAG	180
AAGG'	TGGCCG	ACGCGCTGAC	CAACGCUGTG	GCGCACCTGG	ACGACATGCC	CAACGCCCTG	240
TCCG	ÇÇÇTÜA	GOGACCTGCA	CGCGCACAAG	CTTCGGGTGG	ACCCGGTCAA	CTTCAAGCTC	3.10
CTAA	GCCACT	GCCTGCTGGT	GACCCTGGCC	GCCCACCTCC	CCGCCGAGTI	CACCCCTGCG	340
GTGC.	ACGCCT	CCCTGGACAA	GTTCCTGGCT	TCTGTGAGCA	CCGTGCTGAC	CTCCAAATAC	41.0
CGT							423

# (2) INFORMATION FOR SEQ ID NO:16:

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 141 amino acids
  - (B) TYPE: amino acid
  - (C) STRANDEDNESS:
  - (D) TOPOLOGY: linear
- (ii) MOLECULE TYPE: peptide
- (xi) SEQUENCE DESCRIPTION: SEQ ID NO:16:
- Tal thu Ser Pro Ala Asp Lys Thr Ash Val Lys Ala Ala Trp Gly Lys
- val dry Ala Him Ala Mly dri Tyn dry Ala Him Ala Les His Ard Med Lit
- Fig. 19. Ser Phe Pro Thr Thr Lys Thr Tyr Phe Fro His bhe Asp 2024 35 40 45
- Ser His Gly Ser Ala Gln Val Lys Gly His Gly Lys Lys Val Ala Asp 50
- A, a for The Ash Ala Mal Ala His Mal Asp Asp Mor Pro Ash Ala 100

Leu Ala Ser Van Ser Thr Val Leu Thr Ser Lys Tyr Ard 130 135 140

# (2) INFORMATION FOR SEQ ID NO:10:

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 438 base pairs
  - (B) TYPE: nucleic acid
  - (C) STRANDEINESS: single
  - (D) TOPOLOGY: linear
- (ii) MOLECULE PYPE: DNA (genomic)
- (xi) SEQUENCE DESCRIPTION: SEQ ID NO:17:

GTGCACC	TGA	CTCCTGAGGA	GAAGTCTGCC	GTTACTGCCC	TGTGGGGCAA	GGTGAACGTG	60
GATGAAG	TTG	GTGGTGAGGC	CCTGGGCAGG	CTGCTGGTGG	TCTACCTTTG	GACCCAGAGG	120
TTČTTG	AGT	CCTTTGGGGGA	TCTGTCCACT	CCTGATGCTG	TTATGGGCAA	CCCTAAGGTG	130
AAGGCTC.	ATG	GCAAGAAAGT	GOTOGGTGCC	PTTAGTGATG	GCCTGGCTCA	CCTGGACAAC	240
CTCAAGG	GC.A	CONTROCCAC	ACTGAGTGAG	OTGCACTGT3	ACAAGCIGCA	CGTGGATCCT	300
GAGAACT	TCA	GGCTGCTGGG	CAACGTGCTG	GTCTGTGTGC	TGGCCCATCA	CTTTGGCAAA	360
GAATTCA	CCC	CACCAGTGCA	GGCTGCCTAT	CAGAAAGTGG	TGGCTGGTGT	GGCTAATGCC	420
CTGGCCC.	ACA	AGTATCAC					438

- (E) INFORMATION FOR SEQ ID NO:13:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 146 amino arids
    - (B) TYPE: amino acid
    - (C) STRANDEDNESS:
    - com Topology, linear
  - in MOLECULE TYPE, peptide.
  - wit segrence rescription: Se, 10 N :18:
  - Tall His io. The Bro Will William Son Ala Tall His Alades in the by

Lys Lys Val Leu Gly Ala Phe Ser Asp Gly Leu Ala His Leu Asp Asn 65 70 75 80

Leu Lys Gly Thr Phe Ala Thr Leu Ser Glu Leu His Cys Asp Lys Leu 85 90 95

His Val Asp Pro Glu Asn Phe Arg Leu Leu Gly Asn Val Leu Val Cys
100 105 110

Val Leu Ala His His Phe Gly Lys Glu Phe Thr Pro Pro Val Gln Ala 115 120 125

Ala Tyr Gln Lys Val Val Ala Gly Val Ala Asn Ala Leu Ala His Lys 130 135 140

Tyr His

### (2) INFORMATION FOR SEQ ID NO:19:

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 141 amino acids
  - (B) TYPE: amino acid
  - (C) STRANDEDNESS:
  - (D) TOPOLOGY: linear
- (ii) MOLECULE TYPE: peptide
- (xi) SEQUENCE DESCRIPTION: SEQ ID NO:19:
- Val Leu Ser Gly Glu Asp Lys Ser Asn Ile Lys Ala Ala Trp Gly Lys

  1 10 15
- The Gly Gly His Gly Ala Glu Tyr Gly Ala Glu Ala Leu Glu Arg Met  $10^{\circ}$
- Fig. A.a Ser size Pro Thi Inc. Lys Inc. Ly: Five Fre His Pice Asp Ust.  $4^{\circ}$  .  $4^{\circ}$
- ren Him Mly Jer Ala Min Val Lyr Bly Him Mly Lyr Lyr Val Ala Arr 50 55 60
- Ala Leu Ala Ser Ala Ala Gly His Leu Asp Asp Leu Pro Gly Ala Leu 65 70 75 80
- Ser A.a Len Ser Asp Leu His Ala His Lys Leu Arg Val Asp Pro Val

### (2) INFORMATION FOR SEQ ID NO:20:

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 146 amino acids
  - (B) TYPE: amino acid
  - (C) STRAMDEDNESS:
  - (D) TOPOLOGY: linear
- (ii) MOLECULE TYPE: peptide
- (xi) SEQUENCE DESCRIPTION: SEQ ID NO:20:
- Val His Lei Thr Asp Ala Glu Lys Ala Ala Val Ser Cys Leu Trp Gly
  1 5 10 15
- Lys Val Asn Ser Asp Glu Val Gly Gly Glu Ala Leu Gly Arg Leu Leu 20 25 30
- Val Val Tyr Pro Trp Thr Gln Arg Tyr Phe Asp Ser Phe Gly Asp Leu 35 40 45
- Ser Ser Ala Ser Ala Ile Met Gly Asr. Ala Lys Val Lys Ala His Gly 50 55 60
- Lys Lys Val Ile Thr Ala Phe Asn Asp Gly Leu Asn His Leu Asp Ser 65 70 75 80
- Leu Lys Gly Thr Phe Ala Ser Leu Ser Glu Leu His Cys Asp Lys Leu 85 90 95
- His Val Asp Pro Glu Asn Phe Arg Leu Leu Gly Asn Met Ile Val Ile 100 105 110
- Val Leu Gly His His Leu Gly Lys Asp Phe Thr Pro Ala Ala Gln Ala 115 120 125
- Ala Pho d . Nys Val Val Ala Gly Val Ala The Ala Leu Ala His Nys 130 - 135 - 140
- iye Hib Life
- (1) INFORMATION FOR SEQ ID NO:21:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 141 amino acids
    - .B: TYPE amino acid
    - FOR STRAIDFDNESS:

Val Gly Gly Gln Ala Gly Ala His Gly Ala Glu Ala Leu Glu Arg Met 20 25 30

Phe Lea Gly Phe Pro Thr Thr Lys Thr Tyr Phe Pro His Phe Ash Lea 35 40 45

Ser His Gly Ser Asp Gln Val Lys Ala His Gly Gln Lys Val Ala Asp 50 55 60

Ala Leu Thr Lys Ala Val Gly H.s Leu Asp Asp Leu Pro Gly Ala Leu 65 70 75 80

Ser Ala Leu Ser Asp Leu His Ala His Lys Leu Arg Val Asp Pro Val 85 90 95

Asn Phe Lys Leu Leu Ser His Cys Leu Leu Val Thr Leu Ala Ala His 100 105 110

His Pro Asp Asp Phe Asn Pro Ser Val His Ala Ser Leu Asp Lys Phe 115 120 125

Leu Ala Asn Val Ser Thr Val Leu Thr Ser Lys Tyr Arg 130 135 140

### (2) INFORMATION FOR SEQ ID NO:22:

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 146 amino acids
  - (B) TYPE: amino acid
  - (C) STRANDEDNESS:
  - (D) TOPOLOGY: linear
- (ii) MOLECULE TYPE: peptide

-x.. SEQUENÇE DESCRIPTION: SE, ID Not 20:

yas His ie ser ķīk dit dis iys to Alā Vēl ies ty ies jir ty

typ Val Acc. Val Asp Sin Val Siy Siy Sin Ala Led Siy Arg Leu Let.  $20^\circ$   $-25^\circ$   $-30^\circ$ 

Val Val Tyr Pro Trp Thr Gln Arg Phe Phe Glu Ser Phe Gly Asp Leu 35 40 45

for Ash Ala Ash Ala Val Met Glv Ash Pro Lys Val Lys Ala His Gly

Mal Leu Ala Ard Arg Leu Gly His Asp Pho Ash Pro Asp Mal Glh Ala 120 Ala Phe Gir Lys Val Val Ala Gly Val Ala Ash Ala Leu Ala His Lys 135 Tyr His 145 (2) INFORMATION FOR SEQ ID NO:23: (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 73 amino acids (B) TYPE: amino acid (C) STRANDEDNESS: (D) TOPOLOGY: linear (ii) MOLECULE TYPE: peptide (xi) SEQUENCE DESCRIPTION: SEQ ID NO:23: Val His Let Ser Ala Glu Glu Lys Glu Ala Val Leu Gly Leu Trp Gly Lys Val Ash Val Asp Glu Val (2) INFORMATION FOR SEQ ID NO:24: (1) SEQUENCE CHARACTERISTICS: (A) LENGTH: 20 amino acids (B) TYPE: amino acid (C) STRANDEDNESS: (D) TOPOLOGY: linear \*.i\* M\*C.ECULE CYPE: peptide (xi) SEQUENCE DESCRIPTION: SEG IN NO:24: Val Leu Ser Ala Ala Asp Lys Ala Asp Val Lys Ala Ala Trp Gly Lys 1 5 10 15 mai say Hiy gan 

(ii) MOLECULE TYPE: peptide

(C) STRANDEDNESS:
(D) TOPOLOGY: linear

 $(\pm i)$  SEQUENCE DESCRIPTION: SEQ ID NO:26:

Leu Val Val Tyr Pro Trp Thr Gln Arg Phe 1 5 10

- (2) INFORMATION FOR SEQ ID NO:27:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 8 amino acids
    - (B) TYPE: amino acid
    - (C) STRANDEDNESS:
    - (D) TOPOLOGY: linear
  - (ii) MOLECULE TYPE: peptide
  - RIN SEQUENCE DESCRIPTION: SEA IN NUMBER
  - Tal Tal Tyr Brow Hip Thi Sun Arc
- (L) INFORMATION FOR SEQ ID NO:28:
  - \*ALL SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 4 amino acids
    - (B) TYPE: amino acid

Ty: Pro Trp Thr 1 INFORMATION FOR

### (2) INFORMATION FOR SEQ ID NO:29:

- (i) SEQUÊNCE (HARACTERISTICS:
  - (A) LENGTH: 4 amino acids
  - (B) CYPE: amino acid
  - (C) STRANDEDNESS:
  - (D) TOPOLOGY: linear
- (ii) MOLECULE TYPE: peptide
- (x1) SEQUENCE DESCRIPTION: SEQ ID NO:29:
- Gly Tyr Pro Tyr
- (2) INFORMATION FOR SEQ ID NO:30:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 4 amino acids
    - (B) TYPE: amino acid
    - (C) STRANDEDNESS:
    - (D) TOPOLOGY: linear
  - (ii) MOLECULE TYPE: peptide
  - (x1) SEQUENCE DESCRIPTION: SEQ ID NO:30:
  - Gly Lys Pro Tyr
- . MIFORMATION FOR SEQ ID NO:31:
  - i gegenne e akadikrisilds:
    - (A) LEMWIE: 4 amino acids
    - #F: FYPE amino acil
    - (C) STRANDEDNESS:
    - (D) TOPOLOGY: linear
  - (ii) MOLECULE TYPE: peptide

- (B) TYPE: amino acid
- (C) STRANDEDNESS:
- (D) TOPOLOGY: linear
- (ii) MOLECULE TYPE: peptide
- (xi) SEQUENCE DESCRIPTION: SEQ ID NO:32:

Lys Met Trp Met Phe Arg

- (2) INFORMATION FOR SEQ ID NO:33:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 8 amino acids
    - (E) TYPE: amino arid
    - (C) STRANDEDHESS:
    - (D) TOPOLOGY: linear
  - (ii) MODECULE TYPE: peptide
  - (xi) SEQUENCE DESCRIPTION: SEQ ID NO:33:

Val Arg Arg Met Phe Gly Gly Tyr  $^{4}$ 

- (2) INFORMATION FOR SEQ ID NO:34:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 6 amino acids
    - (B) TYPE: amino acid
    - (C) STRANDEDNESS: single
    - (1) TOPHINGY: linear

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(xi) SEQUENCE DESCRIPTION: SEQ ID No:34:

Phe Leu Gly Phe Pro Thr

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(ix) FEATURE:
         (A) MAME KEY: Modified-site
         (B) LOCATION. 1
         (D) OTHER INFORMATION: product: "position 1"
  label- Xaa
  note: "pyroGln"
   (xi) SEQUENCE DESCRIPTION: SEQ ID NO:35:
    Maa Gln Gln Asp Cys Lys
(2) INFORMATION FOR SEQ ID NO:36:
    (i) SEQUENCE CHARACTERISTICS:
         (A) LENGTH: 4 amino acids
         (B) TYPE: amino acid
         (C) STRANDEDNESS: single
         (D) TOPOLOGY: linear
   (ii) MODECULE TYPE: peptide
   (ix) FEATURE:
         (A) NAME, KEY: Modified-site
         (B) LOCATION: 1
         (D) OTHER INFORMATION: /product= "position 1"
  label= Xaa
  note: "N-AretylSer"
   (xi) SEQUENCE DESCRIPTION: SEQ ID NO:36:
    Xaa Asp Lys Pro
(2) INFORMATION FOR SEQ ID NO:37:

### SEQUENCE CHARACTERISTICS:
         (A) LENGTH: .3 amins acids
         (E) TYPE: am.no acid
         (tt) DIRANDEDHESS: single
         and Top Town Himan
   (ii) MOLECULE TYPE: peptide
    ix FEATURE:
        TANK MAME PEY Modifie i site
         TRU TOCATION: I
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(xi) SEQUENCE DESCRIPTION: SEQ ID NO:37:

Xaa Pro His Xaa Asp Leu Ser His Gly Ser Ala Gln Val